SEP 2 8 2007 20 <1

SEQUENCE LISTING

<110> BROWN, ARTHUR M.
WIBLE, BARBARA A.

<120> METHODS OF INDUCING APOPTOSIS IN HYPERPROLIFERATION CELLS

<130> 22884/04085

<140> 10/784,528

<141> 2004-02-23

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Gly Pro Ser Asp Leu Ser Leu Leu Ser Leu Pro Pro Gly Thr Ser Pro
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gta ggc tcc cct ggt cct cta gct ccc att ccc cca acg ctg ttg gcc 144 Val Gly Ser Pro Gly Pro Leu Ala Pro Ile Pro Pro Thr Leu Leu Ala 35 40 45

cct ggc acc ctg ctg ggc ccc aag cgt gag gtg gac atg cac ccc cct 192
Pro Gly Thr Leu Leu Gly Pro Lys Arg Glu Val Asp Met His Pro Pro

ctg ccc cag cct gtg cac cct gat gtc acc atg aaa cca ttg ccc ttc Leu Pro Gln Pro Val His Pro Asp Val Thr Met Lys Pro Leu Pro Phe
65 70 75 80

tat gaa gtc tat ggg gag ctc atc cgg ccc acc acc ctt gca tcc act

Tyr Glu Val Tyr Gly Glu Leu Ile Arg Pro Thr Thr Leu Ala Ser Thr

tct agc cag cgg ttt gag gaa gcg cac ttt acc ttt gcc ctc aca ccc 336 Ser Ser Gln Arg Phe Glu Glu Ala His Phe Thr Phe Ala Leu Thr Pro 100 105 110

													gga Gly		384
		-			_		_				_		tgt Cys		432
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_			_	_		_	_						acc Thr 175	_	528
				_			_	_		_		_	aca Thr		576
_	_	_		-		_						_	aat Asn		624
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													act Thr 255		768
													ctc Leu		816
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											Leu		atg Met		912
										Asp			gct Ala		960
	_				_				Met				agt Ser 335		1008

	tca Ser			_					-	_	-				_	1056
	atg Met			_	-		_			_	-		_			1104
	999 Gly 370	_	_			-		-		_	_			_		1152
	gag Glu															1200
	tca Ser															1248
	tca Ser	_	_		_	_				_			_	_		1296
	ggc Gly															1344
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gcc Ala 465	ttc Phe	cca Pro	ctg Leu	gga Gly	gcc Ala 470	gac Asp	atc Ile	caa Gln	ggt Gly	tta Leu 475	gat Asp	tta Leu	ttt Phe	tca Ser	ttt Phe 480	1440
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	gaa Glu															1536
	cac His		Leu					Pro					Ser	_		1584
		Thr										Ser			gcc Ala	1632
	-					Glu					Pro				ggt Gly 560	1680

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Val Gly Ser Pro Gly Pro Leu Ala Pro Ile Pro Pro Thr Leu Leu Ala 35 40 45

Pro Gly Thr Leu Leu Gly Pro Lys Arg Glu Val Asp Met His Pro Pro 50 55 60

Leu Pro Gln Pro Val His Pro Asp Val Thr Met Lys Pro Leu Pro Phe 65 70 75 80

Tyr Glu Val Tyr Gly Glu Leu Ile Arg Pro Thr Thr Leu Ala Ser Thr 85 90 95

Ser Ser Gln Arg Phe Glu Glu Ala His Phe Thr Phe Ala Leu Thr Pro 100 105 110

Gln Gln Val Gln Gln Ile Leu Thr Ser Arg Glu Val Leu Pro Gly Ala 115 120 125

Lys Cys Asp Tyr Thr Ile Gln Val Gln Leu Arg Phe Cys Leu Cys Glu 130 135 140

Thr Ser Cys Pro Gln Glu Asp Tyr Phe Pro Pro Asn Leu Phe Val Lys 145 150 155 160

Val Asn Gly Lys Leu Cys Pro Leu Pro Gly Tyr Leu Pro Pro Thr Lys 165 170 175

Asn Gly Ala Glu Pro Lys Arg Pro Ser Arg Pro Ile Asn Ile Thr Pro 180 185 190

Leu Ala Arg Leu Ser Ala Thr Val Pro Asn Thr Ile Val Val Asn Trp
195 200 205

Ser Ser Glu Phe Gly Arg Asn Tyr Ser Leu Ser Val Tyr Leu Val Arg 210 215 220

Gln Leu Thr Ala Gly Thr Leu Leu Gln Lys Leu Arg Ala Lys Gly Ile 225 230 235 240

Arg Asn Pro Asp His Ser Arg Ala Leu Ile Lys Glu Lys Leu Thr Ala 245 250 255

- Asp Pro Asp Ser Glu Val Ala Thr Thr Ser Leu Arg Val Ser Leu Met 260 265 270
- Cys Pro Leu Gly Lys Met Arg Leu Thr Val Pro Cys Arg Ala Leu Thr 275 280 285
- Cys Ala His Leu Gln Ser Phe Asp Ala Ala Leu Tyr Leu Gln Met Asn 290 295 300
- Glu Lys Lys Pro Thr Trp Thr Cys Pro Val Cys Asp Lys Lys Ala Pro 305 310 315 320
- Tyr Glu Ser Leu Ile Ile Asp Gly Leu Phe Met Glu Ile Leu Ser Ser 325 330 335
- Cys Ser Asp Cys Asp Glu Ile Gln Phe Met Glu Asp Gly Ser Trp Cys 340 345 350
- Pro Met Lys Pro Lys Lys Glu Ala Ser Glu Val Cys Pro Pro Gly 355 360 365
- Tyr Gly Leu Asp Gly Leu Gln Tyr Ser Pro Val Gln Gly Gly Asp Pro 370 375 380
- Ser Glu Asn Lys Lys Lys Val Glu Val Ile Asp Leu Thr Ile Glu Ser 385 390 395 400
- Ser Ser Asp Glu Glu Asp Leu Pro Pro Thr Lys Lys His Cys Ser Val 405 410 415
- Thr Ser Ala Ala Ile Pro Ala Leu Pro Gly Ser Lys Gly Val Leu Thr
 420 425 430
- Ser Gly His Gln Pro Ser Ser Val Leu Arg Ser Pro Ala Met Gly Thr 435 440 445
- Leu Gly Gly Asp Phe Leu Ser Ser Leu Pro Leu His Glu Tyr Pro Pro 450 455 460
- Ala Phe Pro Leu Gly Ala Asp Ile Gln Gly Leu Asp Leu Phe Ser Phe 465 470 475 480
- Leu Gln Thr Glu Ser Gln His Tyr Gly Pro Ser Val Ile Thr Ser Leu 485 490 495
- Asp Glu Gln Asp Ala Leu Gly His Phe Phe Gln Tyr Arg Gly Thr Pro 500 505 510
- Ser His Phe Leu Gly Pro Leu Ala Pro Thr Leu Gly Ser Ser His Cys 515 520 525
- Ser Ala Thr Pro Ala Pro Pro Pro Gly Arg Val Ser Ser Ile Val Ala 530 · 540
- Pro Gly Gly Ala Leu Arg Glu Gly His Gly Gly Pro Leu Pro Ser Gly 545 550 555

Pro Ser Leu Thr Gly Cys Arg Ser Asp Ile Ile Ser Leu Asp 565 570

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Thr Leu Leu Gly Pro Lys Arg Glu Val Asp Met His
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<210> 4
<211> 4
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<213> Artificial Sequence
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      peptide
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Asp Glu Val Asp
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<213> Homo sapiens
<400> 5
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Ser Glu Leu Gln Val Leu Leu Gly Tyr Ala Gly Arg Asn Lys His Gly
Arg Lys His Glu Leu Leu Thr Lys Ala Leu His Leu Leu Lys Ala Gly
Cys Ser Pro Ala Val Gln Met Lys Ile Lys Glu Leu Tyr Arg Arg Arg
Phe Pro Gln Lys Ile Met Thr Pro Ala Asp Leu Ser Ile Pro Asn Val
                                          75
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His Ser Ser Pro Met Pro Ala Thr Leu Ser Pro Ser Thr Ile Pro Gln 85 90 95

man in the relies in a militaritation will be the contraction

- Leu Thr Tyr Asp Gly His Pro Ala Ser Ser Pro Leu Leu Pro Val Ser 100 105 110
- Leu Leu Gly Pro Lys His Lys Leu Glu Leu Pro His Leu Thr Ser Ala 115 120 125
- Leu His Pro Val His Pro Asp Ile Lys Leu Gln Lys Leu Pro Phe Tyr 130 135 140
- Asp Leu Leu Asp Glu Leu Ile Lys Pro Thr Ser Leu Ala Ser Asp Asn 145 150 155 160
- Ser Gln Arg Phe Arg Glu Thr Cys Phe Ala Phe Ala Leu Thr Pro Gln
 165 170 175
- Gln Val Gln Gln Ile Ser Ser Ser Met Asp Ile Ser Gly Thr Lys Cys 180 185 190
- Asp Phe Thr Val Gln Val Gln Leu Arg Phe Cys Leu Ser Glu Thr Ser 195 200 205
- Cys Pro Gln Glu Asp His Phe Pro Pro Asn Leu Cys Val Lys Val Asn 210 215 220
- Thr Lys Pro Cys Ser Leu Pro Gly Tyr Leu Pro Pro Thr Lys Asn Gly 225 230 235 240
- Val Glu Pro Lys Arg Pro Ser Arg Pro Ile Asn Ile Thr Ser Leu Val 245 250 255
- Arg Leu Ser Thr Thr Val Pro Asn Thr Met Cys Ser Trp Thr Ala Glu 260 265 270
- Ile Gly Arg Asn Tyr Ser Met Ala Val Tyr Leu Val Lys Gln Leu Ser 275 280 285
- Ser Thr Val Leu Gln Arg Leu Arg Ala Lys Gly Ile Arg Asn Pro 290 295 300
- Asp His Ser Arg Ala Leu Ile Lys Glu Lys Leu Thr Ala Asp Pro Asp 305 310 315 320
- Ser Glu Ile Ala Thr Thr Ser Leu Arg Val Ser Leu Leu Cys Pro Leu 325 330 335
- Gly Lys Met Arg Leu Thr Ile Pro Cys Arg Ala Leu Thr Cys Ser His \$340\$ \$350 $^{\circ}$
- Leu Gln Cys Phe Asp Ala Thr Leu Tyr Ile Gln Met Asn Glu Lys Lys \cdot 355 360 365
- Pro Thr Trp Val Cys Pro Val Cys Asp Lys Lys Ala Pro Tyr Glu His

Leu Ile Ile Asp Gly Leu Phe Met Glu Ile Leu Lys Tyr Cys Thr Asp 385 390 395 400

Cys Asp Glu Ile Gln Phe Lys Glu Asp Gly Thr Trp Ala Pro Met Arg
405 410 415

Ser Lys Lys Glu Val Gln Glu Val Ser Ala Ser Tyr Asn Gly Val Asp 420 425 430

Gly Cys Leu Ser Ser Thr Leu Glu His Gln Val Ala Ser His His Gln 435 440 445

Ser Ser Asn Lys Asn Lys Lys Val Glu Val Ile Asp Leu Thr Ile Asp 450 450

Ser Ser Ser Asp Glu Glu Glu Glu Fro Ser Ala Lys Arg Thr Cys 475 480

Pro Ser Leu Ser Pro Thr Ser Pro Leu Asn Asn Lys Gly Ile Leu Ser 485 490 495

Leu Pro His Gln Ala Ser Pro Val Ser Arg Thr Pro Ser Leu Pro Ala 500 . 505 510

Val Asp Thr Ser Tyr Ile Asn Thr Ser Leu Ile Gln Asp Tyr Arg His 515 520 525

Pro Phe His Met Thr Pro Met Pro Tyr Asp Leu Gln Gly Leu Asp Phe 530 540

Phe Pro Phe Leu Ser Gly Asp Asn Gln His Tyr Asn Thr Ser Leu Leu 545 550 555

Ala Ala Ala Ala Ala Val Ser Asp Gln Asp Leu Leu His Ser 565 570 575

Ser Arg Phe Phe Pro Tyr Thr Ser Ser Gln Met Phe Leu Asp Gln Leu 580 585 590

Ser Ala Gly Gly Ser Thr Ser Leu Pro Thr Thr Asn Gly Ser Ser Ser 595 600 605

Gly Ser Asn Ser Ser Leu Val Ser Ser Asn Ser Leu Arg Glu Ser His 610 615 620

Ser His Thr Val Thr Asn Arg Ser Ser Thr Asp Thr Ala Ser Ile Phe 625 630 635 640

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<211> 651

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Ser Glu Leu Gln Val Leu Leu Gly Tyr Ala Gly Arg Asn Lys His Gly 20 25 30

Arg Lys His Glu Leu Leu Thr Lys Ala Leu His Leu Leu Lys Ala Gly 35 40 45

Cys Ser Pro Ala Val Gln Met Lys Ile Lys Glu Leu Tyr Arg Arg 50 55 60

Phe Pro Gln Lys Ile Met Thr Pro Ala Asp Leu Ser Ile Pro Asn Val 65 70 75 80

His Ser Ser Pro Met Pro Pro Thr Leu Ser Pro Ser Thr Ile Pro Gln
85 90 95

Leu Thr Tyr Asp Gly His Pro Ala Ser Ser Pro Leu Leu Pro Val Ser 100 105 110

Leu Leu Gly Pro Lys His Glu Leu Glu Leu Pro His Leu Thr Ser Ala 115 120 125

Leu His Pro Val His Pro Asp Ile Lys Leu Gln Lys Leu Pro Phe Tyr 130 135 140

Asp Leu Leu Asp Glu Leu Ile Lys Pro Thr Ser Leu Ala Ser Asp Asn 145 150 155 160

Ser Gln Arg Phe Arg Glu Thr Cys Phe Ala Phe Ala Leu Thr Pro Gln 165 170 175

Gln Val Gln Gln Ile Ser Ser Ser Met Asp Ile Ser Gly Thr Lys Cys 180 185 190

Asp Phe Thr Val Gln Val Gln Leu Arg Phe Cys Leu Ser Glu Thr Ser 195 200 205

Cys Pro Gln Glu Asp His Phe Pro Pro Asn Leu Cys Val Lys Val Asn 210 215 220

Thr Lys Pro Cys Ser Leu Pro Gly Tyr Leu Pro Pro Thr Lys Asn Gly 225 230 235

Val Glu Pro Lys Arg Pro Ser Arg Pro Ile Asn Ile Thr Ser Leu Val 245 250 255

Arg Leu Ser Thr Thr Val Pro Asn Thr Ile Val Val Ser Trp Thr Ala 260 265 270

Glu Ile Gly Arg Asn Tyr Ser Met Ala Val Tyr Leu Val Lys Gln Leu 275 280 285

Ser Ser Thr Val Leu Leu Gln Arg Leu Arg Ala Lys Gly Ile Arg Asn 290 295 300

- Pro Asp His Ser Arg Ala Leu Ile Lys Glu Lys Leu Thr Ala Asp Ser 305 310 315 320
- Asp Ser Glu Ile Ala Thr Thr Ser Leu Arg Val Ser Leu Leu Cys Pro 325 330 335
- Leu Gly Lys Met Arg Leu Thr Ile Pro Cys Arg Ala Leu Thr Cys Ser 340 345 350
- His Leu Gln Cys Phe Asp Ala Thr Leu Tyr Ile Gln Met Asn Glu Lys 355 360 365
- Lys Pro Thr Trp Val Cys Pro Val Cys Asp Lys Lys Ala Pro Tyr Glu 370 375 380
- His Leu Ile Ile Asp Gly Leu Phe Met Glu Ile Leu Lys Tyr Cys Thr 385 390 395 400
- Asp Cys Asp Glu Ile Gln Phe Lys Glu Asp Gly Ser Trp Ala Pro Met
 405 410 415
- Arg Ser Lys Lys Glu Val Gln Glu Val Thr Ala Ser Tyr Asn Gly Val 420 425 430
- Asp Gly Cys Leu Ser Ser Thr Leu Glu His Gln Val Ala Ser His Asn
 435
 440
 445
- Gln Ser Ser Asn Lys Asn Lys Lys Val Glu Val Ile Asp Leu Thr Ile 450 455 460
- Asp Ser Ser Ser Asp Glu Glu Glu Glu Pro Pro Ala Lys Arg Thr 465 470 475 480
- Cys Pro Ser Leu Ser Pro Thr Ser Pro Leu Ser Asn Lys Gly Ile Leu 485 490 495
- Ser Leu Pro His Gln Ala Ser Pro Val Ser Arg Thr Pro Ser Leu Pro 500 505 510
- Ala Val Asp Thr Ser Tyr Ile Asn Thr Ser Leu Ile Gln Asp Tyr Arg 515 520 525
- His Pro Phe His Met Thr Pro Met Pro Tyr Asp Leu Gln Gly Leu Asp 530 535 540
- Phe Phe Pro Phe Leu Ser Gly Asp Asn Gln His Tyr Asn Thr Ser Leu 545 550 555 560
- Leu Ala Ala Ala Ala Ala Val Ser Asp Asp Gln Asp Leu Leu His
 565 570 575
- Ser Ser Arg Phe Phe Pro Tyr Thr Ser Ser Gln Met Phe Leu Asp Gln 580 585 590
- Leu Ser Ala Gly Gly Ser Thr Ser Leu Pro Ala Thr Asn Gly Ser Ser 595 600 605

Ser Gly Ser Asn Ser Ser Leu Val Ser Ser Asn Ser Leu Arg Glu Ser 610 620

His Gly His Gly Val Ala Ser Arg Ser Ser Ala Asp Thr Ala Ser Ile 625 630 635 640

Phe Gly Ile Ile Pro Asp Ile Ile Ser Leu Asp 645 650

<210> 7

<211> 583

<212> PRT

<213> Mus musculus

<400> 7

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Ala Gly Arg Asn Lys Ser Gly Arg Lys His Glu Leu Leu Ala Lys Ala 20 25 30

Leu His Leu Leu Lys Ser Ser Cys Ala Pro Ser Val Gln Met Lys Ile 35 40 45

Lys Glu Leu Tyr Arg Arg Phe Pro Arg Lys Thr Leu Gly Pro Ser
50 55 60

Asp Leu Ser Leu Leu Ser Leu Pro Pro Gly Thr Ser Pro Pro Val His 65 70 75 80

Pro Asp Val Thr Met Lys Pro Leu Pro Phe Tyr Glu Val Tyr Gly Glu 85 90 95

Leu Ile Arg Pro Thr Thr Leu Ala Ser Thr Ser Ser Gln Arg Phe Glu 100 105 110

Glu Ala His Phe Thr Phe Ala Leu Thr Pro Gln Gln Leu Gln Gln Ile 115 120 125

Leu Thr Ser Arg Glu Val Leu Pro Gly Ala Lys Cys Asp Tyr Thr Ile 130 135 140

Gln Val Gln Leu Arg Phe Cys Leu Cys Glu Thr Ser Cys Pro Gln Glu 145 150 155 160

Asp Tyr Phe Pro Pro Asn Leu Phe Val Lys Val Asn Gly Lys Leu Cys 165 170 175

Pro Leu Pro Gly Tyr Leu Pro Pro Thr Lys Asn Gly Ala Glu Pro Arg 180 185 190

Gly Pro Ala Val Arg Ser Thr Ser His Pro Trp Leu Asp Ser Gln Pro
195 200 205

Leu Ser Pro Thr Pro Ser Leu Leu Ile Gly His Leu Ser Leu Asp Gly 210 215 220

Ile Thr Pro Cys Pro Cys Leu Val Arg Gln Leu Thr Ala Gly Thr Leu 225 230 235 240

. A ministration of the state o

- Leu Gln Lys Leu Arg Ala Lys Gly Ile Arg Asn Pro Asp His Ser Arg
 245 250 255
- Ala Leu Ile Lys Glu Lys Leu Thr Ala Asp Pro Asp Ser Glu Val Ala 260 265 270
- Thr Thr Ser Leu Pro Gly Val Thr His Val Pro Ala Arg Lys Met Arg 275 280 285
- Leu Thr Val Pro Cys Arg Ala Leu Thr Cys Ala His Leu Gln Ser Phe 290 295 300
- Asp Ala Ala Leu Tyr Ile Gln Met Asn Glu Lys Lys Pro Thr Trp Thr 305 310 315 320
- Cys Pro Val Cys Asp Lys Lys Ala Pro Tyr Glu Ser Leu Ile Ile Asp 325 330 335
- Gly Leu Phe Met Glu Ile Leu Asn Ser Cys Ser Asp Cys Asp Glu Ile 340 345 350
- Gln Phe Met Glu Asp Gly Ser Trp Cys Pro Met Lys Pro Lys Lys Glu 355 360 365
- Ala Ser Glu Val Cys Pro Pro Pro Gly Tyr Gly Leu Asp Gly Leu Gln 370 380
- Tyr Ser Ala Val Gln Glu Gly Ile Gln Pro Glu Ser Lys Lys Arg Val
- Glu Val Ile Asp Leu Thr Ile Glu Ser Ser Ser Asp Glu Glu Asp Leu
 405 410 415
- Pro Pro Thr Lys Lys Gln Cys Ser Val Thr Ser Ala Ala Ile Pro Ala 420 425 430
- Leu Leu Gly Ser Lys Gly Val Leu Thr Ser Gly His Gln Pro Ser Ser 435 440 445
- Val Leu Arg Ser Pro Ala Met Gly Thr Leu Gly Ser Asp Phe Leu Ser 450 455 460
- Ser Leu Pro Val His Glu Tyr Pro Pro Ala Phe Pro Leu Gly Ala Asp 465 470 475 480
- Ile Gln Gly Leu Asp Leu Phe Ser Phe Leu Gln Thr Glu Ser Gln Gln 485 490 495
- Tyr Gly Pro Ser Val Ile Ile Ser Leu Asp Glu Gln Asp Thr Leu Gly 500 505 510
- His Pro Phe Gln Tyr Arg Gly Thr Pro Ser His Phe Leu Gly Pro Leu 515 520 525

Ala Pro Thr Leu Gly Ser Cys His Gly Ser Ser Thr Pro Ala Pro Pro 530 540

Pro Gly Arg Val Ser Ser Ile Val Ala Pro Gly Ser Ser Leu Arg Glu 545 550 560

Gly His Gly Gly Pro Leu Pro Ser Gly Pro Ser Leu Thr Gly Cys Arg 565 570 575

Ser Asp Val Ile Ser Leu Asp 580

A series of collections of the series of the

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<211> 572

<212> PRT

<213> Homo sapiens

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Ser Glu Leu Gln Val Leu Gly Phe Ala Gly Arg Asn Lys Ser Gly 20 25 30

Arg Lys His Asp Leu Leu Met Arg Ala Leu His Leu Leu Lys Ser Gly
35 40 45

Cys Ser Pro Ala Val Gln Ile Lys Ile Arg Glu Leu Tyr Arg Arg Arg 50 55 60

Tyr Pro Arg Thr Leu Glu Gly Leu Ser Asp Leu Ser Thr Ile Lys Ser 65 70 75 80

Ser Val Phe Ser Leu Asp Gly Gly Ser Ser Pro Val Glu Pro Asp Leu 85 90 95

Ala Val Ala Gly Ile His Ser Leu Pro Ser Thr Ser Val Thr Pro His

Ser Pro Ser Ser Pro Val Gly Ser Val Leu Leu Gln Asp Thr Lys Pro 115 120 125

Thr Phe Glu Met Gln Gln Pro Ser Pro Pro Ile Pro Pro Val His Pro 130 135 140

Asp Val Gln Leu Lys Asn Leu Pro Phe Tyr Asp Val Leu Asp Val Leu 145 150 155 160

Ile Lys Pro Thr Ser Leu Val Gln Ser Ser Ile Gln Arg Phe Gln Glu 165 170 175

Lys Phe Phe Ile Phe Ala Leu Thr Pro Gln Gln Val Arg Glu Ile Cys 180 185 190

Ile Ser Arg Asp Phe Leu Pro Gly Gly Arg Arg Asp Tyr Thr Val Gln 195 200 205

Val Gln Leu Arg Leu Cys Leu Ala Glu Thr Ser Cys Pro Gln Glu Asp 210 215 220

A STATE OF THE PROPERTY OF THE

- Asn Tyr Pro Asn Ser Leu Cys Ile Lys Val Asn Gly Lys Leu Phe Pro 225 230 235 240
- Leu Pro Gly Tyr Ala Pro Pro Pro Lys Asn Gly Ile Glu Gln Lys Arg
 245 250 255
- Pro Gly Arg Pro Leu Asn Ile Thr Ser Leu Val Arg Leu Ser Ser Ala 260 265 270
- Val Pro Asn Gln Ile Ser Ile Ser Trp Ala Ser Glu Ile Gly Lys Asn 275 280 285
- Tyr Ser Met Ser Val Tyr Leu Val Arg Gln Leu Thr Ser Ala Met Leu 290 295 300
- Leu Gln Arg Leu Lys Met Lys Gly Ile Arg Asn Pro Asp His Ser Arg 305 310 315 320
- Ala Leu Ile Lys Glu Lys Leu Thr Ala Asp Pro Asp Ser Glu Ile Ala 325 330 335
- Thr Thr Ser Leu Arg Val Ser Leu Met Cys Pro Leu Gly Lys Met Arg
- Leu Thr Ile Pro Cys Arg Ala Val Thr Cys Thr His Leu Gln Cys Phe 355 360 365
- Asp Ala Ala Leu Tyr Ile Gln Met Asn Glu Lys Lys Pro Thr Trp Ile 370 375 380
- Cys Pro Val Cys Asp Lys Lys Ala Ala Tyr Glu Ser Leu Ile Leu Asp 385 390 395 400
- Gly Leu Phe Met Glu Ile Leu Asn Asp Cys Ser Asp Val Asp Glu Ile 405 410 415
- Lys Phe Gln Glu Asp Gly Ser Trp Cys Pro Met Arg Pro Lys Lys Glu 420 425 430
- Ala Met Lys Val Ser Ser Gln Pro Cys Thr Lys Ile Glu Ser Ser Ser 435 440 445
- Val Leu Ser Lys Pro Cys Ser Val Thr Val Ala Ser Glu Ala Ser Lys 450 455 460
- Lys Lys Val Asp Val Ile Asp Leu Thr Ile Glu Ser Ser Asp Glu 465 470 475 480
- Glu Glu Asp Pro Pro Ala Lys Arg Lys Cys Ile Phe Met Ser Glu Thr \$485\$
- Gln Ser Ser Pro Thr Lys Gly Val Leu Met Tyr Gln Pro Ser Ser Val 500 505

Arg Val Pro Ser Val Thr Ser Val Asp Pro Ala Ala Ile Pro Pro Ser 515 520 525

Leu Thr Asp Tyr Ser Val Pro Phe His His Thr Pro Ile Ser Ser Met 530 540

Ser Ser Asp Leu Pro Gly Glu Gln Arg Phe Asn Asp Ile Asn Asn Glu 545 550 560

Leu Lys Leu Gly Thr Ser Ser Asp Thr Val Gln Gln 565 570

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<212> PRT

<213> Homo sapiens

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Met Ala Asp Phe Glu Glu Leu Arg Asn Met Val Ser Ser Phe Arg Val 1 5 10 15

Ser Glu Leu Gln Val Leu Leu Gly Phe Ala Gly Arg Asn Lys Ser Gly 20 25 30

Arg Lys His Asp Leu Leu Met Arg Ala Leu His Leu Leu Lys Ser Gly
35 40 45

Cys Ser Pro Ala Val Gln Ile Lys Ile Arg Glu Leu Tyr Arg Arg Arg 50 55 60

Tyr Pro Arg Thr Leu Glu Gly Leu Ser Asp Leu Ser Thr Ile Lys Ser 65 70 75 80

Ser Val Phe Ser Leu Asp Gly Gly Ser Ser Pro Val Glu Pro Asp Leu 85 90 95

Ala Val Ala Gly Ile His Ser Leu Pro Ser Thr Ser Val Thr Pro His

Ser Pro Ser Ser Pro Val Gly Ser Val Leu Leu Gln Asp Thr Lys Pro 115 120 125

Thr Phe Glu Met Gln Gln Pro Ser Pro Pro Ile Pro Pro Val His Pro 130 135 140

Ile Lys Pro Thr Ser Leu Val Gln Ser Ser Ile Gln Arg Phe Gln Glu 165 170 175

Lys Phe Phe Ile Phe Ala Leu Thr Pro Gln Gln Val Arg Glu Ile Cys 180 185 190

Ile Ser Arg Asp Phe Leu Pro Gly Gly Arg Arg Asp Tyr Thr Val Gln
195 200 205

Val Gln Leu Arg Leu Cys Leu Ala Glu Thr Ser Cys Pro Gln Glu Asp 210 215 220

at all a transfer of the state of the state

- Asn Tyr Pro Asn Ser Leu Cys Ile Lys Val Asn Gly Lys Leu Phe Pro 225 230 235 240
- Leu Pro Gly Tyr Ala Pro Pro Pro Lys Asn Gly Ile Glu Gln Lys Arg
 245 250 255
- Pro Gly Arg Pro Leu Asn Ile Thr Ser Leu Val Arg Leu Ser Ser Ala 260 265 270
- Val Pro Asn Gln Ile Ser Ile Ser Trp Ala Ser Glu Ile Gly Lys Asn 275 280 285
- Tyr Ser Met Ser Val Tyr Leu Val Arg Gln Leu Thr Ser Ala Met Leu 290 295 300
- Leu Gln Arg Leu Lys Met Lys Gly Ile Arg Asn Pro Asp His Ser Arg 305 310 315 320
- Ala Leu Ile Lys Glu Lys Leu Thr Ala Asp Pro Asp Ser Glu Ile Ala 325 330 335
- Thr Thr Ser Leu Arg Val Ser Leu Met Cys Pro Leu Gly Lys Met Arg
- Leu Thr Ile Pro Cys Arg Ala Val Thr Cys Thr His Leu Gln Cys Phe 355 360 365
- Asp Ala Ala Leu Tyr Ile Gln Met Asn Glu Lys Lys Pro Thr Trp Ile 370 375 380
- Cys Pro Val Cys Asp Lys Lys Ala Ala Tyr Glu Ser Leu Ile Leu Asp 385 390 395 400
- Gly Leu Phe Met Glu Ile Leu Asn Asp Cys Ser Asp Val Asp Glu Ile 405 410 415
- Lys Phe Gln Glu Asp Gly Ser Trp Cys Pro Met Arg Pro Lys Lys Glu 420 425 430
- Ala Met Lys Val Ser Ser Gln Pro Cys Thr Lys Ile Glu Ser Ser Ser 435 440 445
- Val Leu Ser Lys Pro Cys Ser Val Thr Val Ala Ser Glu Ala Ser Lys 450 455 460
- Lys Lys Val Asp Val Ile Asp Leu Thr Ile Glu Ser Ser Ser Asp Glu 465 470 475 480
- Glu Glu Asp Pro Pro Ala Lys Arg Lys Cys Ile Phe Met Ser Glu Thr \$485\$
- Gln Ser Ser Pro Thr Lys Gly Val Leu Met Tyr Gln Pro Ser Ser Val 500 505 510

Arg Val Pro Ser Val Thr Ser Val Asp Pro Ala Ala Ile Pro Pro Ser 515 520 525

Leu Thr Asp Tyr Ser Val Pro Phe His His Thr Pro Ile Ser Ser Met 530 540

Ser Ser Asp Leu Pro Gly Leu Asp Phe Leu Ser Leu Ile Pro Val Asp 545 550 555 560

Pro Gln Tyr Cys Pro Pro Met Phe Leu Asp Ser Leu Thr Ser Pro Leu 565 570 575

Thr Ala Ser Ser Thr Ser Val Thr Thr Thr Ser Ser His Glu Ser Ser 580 585 590

Thr His Val Ser Ser Ser Ser Ser Arg Ser Glu Thr Gly Val Ile Thr 595 600 605

Ser Ser Gly Ser Asn Ile Pro Glu Ile Ile Ser Leu Asp 610 615 620

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<212> PRT

<213> Homo sapiens

<400> 10

Met Ala Ala Glu Leu Val Glu Ala Lys Asn Met Val Met Ser Phe Arg 1 5 10 15

Val Ser Asp Leu Gln Met Leu Leu Gly Phe Val Gly Arg Ser Lys Ser 20 25 30

Gly Leu Lys His Glu Leu Val Thr Arg Ala Leu Gln Leu Val Gln Pro 35 40 45

Asp Cys Ser Pro Glu Leu Phe Lys Lys Ile Lys Glu Leu Tyr Glu Thr 50 55 60

Arg Tyr Ala Lys Lys Asn Ser Glu Pro Ala Pro Gln Pro His Arg Pro 65 70 75 80

Leu Asp Pro Leu Thr Met His Ser Thr Tyr Asp Arg Ala Gly Ala Val 85 90 95

Pro Arg Thr Pro Leu Ala Gly Phe Asn Ile Asp Tyr Pro Val Leu Tyr
100 105 110

Gly Lys Tyr Leu Asn Gly Leu Gly Arg Leu Pro Ala Lys Thr Leu Lys 115 120 125

Pro Glu Val Arg Leu Val Lys Leu Pro Phe Phe Asn Met Leu Asp Glu 130 135 140

Glu Ser Pro Cys Ile Phe Ala Leu Thr Pro Arg Gln Val Glu Leu Ile 165 170 175

and a second resident and a second se

- Arg Phe Lys Gln Gly Met Gln Pro Gly Val Lys Ala Val Gln Val Val 180 \$185
- Leu Arg Ile Cys Tyr Ser Asp Thr Ser Cys Pro Gln Glu Asp Gln Tyr 195 200 205
- Pro Pro Asn Ile Ala Val Lys Val Asn His Ser Tyr Cys Ser Val Pro 210 215 220
- Gly Tyr Tyr Pro Ser Asn Lys Pro Gly Val Glu Pro Lys Arg Pro Cys 225 230 235 240
- Arg Pro Ile Asn Leu Thr His Leu Met Tyr Leu Ser Ser Ala Thr Asn 245 250 255
- Arg Ile Thr Val Thr Trp Gly Asn Tyr Gly Lys Ser Tyr Ser Val Ala 260 265 270
- Leu Tyr Leu Val Arg Gln Leu Thr Ser Ser Glu Leu Leu Gln Arg Leu 275 280 285
- Lys Thr Ile Gly Val Lys His Pro Glu Leu Cys Lys Ala Leu Val Lys 290 295 300
- Glu Lys Leu Arg Leu Asp Pro Asp Ser Glu Ile Ala Thr Thr Gly Val 305 310 315 320
- Arg Val Ser Leu Ile Cys Pro Leu Val Lys Met Arg Leu Ser Val Pro 325 330 335
- Cys Arg Ala Glu Thr Cys Ala His Leu Gln Cys Phe Asp Ala Val Phe 340 345 350
- Tyr Ile Gln Met Asn Glu Lys Lys Pro Thr Trp Met Cys Pro Val Cys 355 360 365
- Asp Lys Pro Ala Pro Tyr Asp Gln Leu Ile Ile Asp Gly Leu Leu Ser 370 375 380
- Lys Ile Leu Ser Glu Cys Glu Asp Ala Asp Glu Ile Glu Tyr Leu Val 385 390 395 400
- Asp Gly Ser Trp Cys Pro Ile Arg Ala Glu Lys Glu Arg Ser Cys Ser 405 410 415
- Pro Gln Gly Ala Ile Leu Val Leu Gly Pro Ser Asp Ala Asn Gly Leu 420 425 430
- Leu Pro Ala Pro Ser Val Asn Gly Ser Gly Ala Leu Gly Ser Thr Gly 435 440 445
- Gly Gly Gly Pro Val Gly Ser Met Glu Asn Gly Lys Pro Gly Ala Asp 450 460

Val Val Asp Leu Thr Leu Asp Ser Ser Ser Ser Ser Glu Asp Glu Glu 465 470 475 480

Glu Glu Glu Glu Glu Glu Asp Glu Asp Glu Glu Gly Pro Arg Pro 485 490 495

Lys Arg Arg Cys Pro Phe Gln Lys Gly Leu Val Pro Ala Cys 500 505 505